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ring nodes :
1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20
chain bonds :
1-23 7-24 13-22 15-30 18-29 20-21 23-25 24-26 25-27 26-27 27-28 29-31
30-32
ring bonds :
1-2 1-6 2-3 2-13 3-4 3-16 4-5 5-6 7-8 7-12 8-9 9-10 10-11 11-12 11-17
12-20 13-14 14-15 15-16 17-18 18-19 19-20
exact/norm bonds :
1-23 7-24 13-22 20-21 23-25 24-26 27-28 29-31 30-32
exact bonds :
2-13 3-16 11-17 12-20 13-14 14-15 15-16 15-30 17-18 18-19 18-29 19-20
25-27 26-27
normalized bonds :
1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-12 8-9 9-10 10-11 11-12
isolated ring systems :
containing 1 : 7 :
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Match level :

chain nodes :

21 22 23 24 25 26 27 28 29 30 31 32

 1:CLASS
 2:CLASS
 3:CLASS
 4:CLASS
 5:CLASS
 6:CLASS
 7:CLASS
 8:CLASS
 9:CLASS

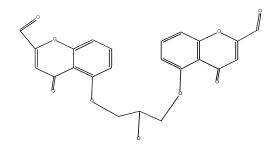
 10:CLASS
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L1 STRUCTURE UPLOADED

=> d 11 L1 HAS NO ANSWERS L1 STR



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146 TO

694

=> s 11

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SAMPLE SCREEN SEARCH COMPLETED - 24 TO ITERATE

100.0% PROCESSED 24 ITERATIONS

SEARCH TIME: 00.00.01

21 ANSWERS

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 187 TO 773

L2 21 SEA SSS SAM L1

=> s 11 ful

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FULL SCREEN SEARCH COMPLETED - 668 TO ITERATE

100.0% PROCESSED 668 ITERATIONS

530 ANSWERS

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=> s 13
L4
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.4 2459 L3

=> d abafbib hitstre 2440-2459

'ABAFBIB' IS NOT A VALID FORMAT FOR FILE 'CAPLUS'

'HITSTRE' IS NOT A VALID FORMAT FOR FILE 'CAPLUS'

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T. 4

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L1 STRUCTURE UPLOADED

L2 21 S L1

L3 530 S L1 FUL

FILE 'CAPLUS' ENTERED AT 10:51:09 ON 30 JUN 2010 2459 S L3

=> d 14 abs fbib hitstr 2440-2459

4 ANSWER 2440 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN

GI For diagram(s), see printed CA Issue.

The title compds. (I), in which X is a polymethylene or a hydroxypolymethylene, and R is H, HO, alkoxy hydroxyalkoxy, or alkenyl, are described. I are useful in the treatment of allergic phenomena such as asthma, hayfever, urticaria, and auto-immune diseases, and they augment the action of antisera. They were prepared by first joining 2 moles of a chromone precursor by the group X to form II, then forming the 4-pyrone ring in II to form I. A mixture of 2,6-dihydroxy-4-methoxyacetophenone (III) and epichlorohydrin was added to a solution of BtONa in BtOH and the mixture refluxed for 4.5 hr to yield II(OXO = 3-OCH2CH(OH)CH2O-3', R = 5-MeO) (IV). In another exp., a mixture of Br(CH2)5Er, III, K2CO3, a trace KI, and Me2CO was refluxed for 4 days to give II (OXO = 3-O(CH2)5-O-3', R = 5-MeO). A solution of IV and (COZEt)2 in dioxame-EtOH was added to a solution of EtONa in EtOH and the mixture refluxed for 4 hr to yield I (OXO = 5-OCH2CH(OH)CH2O-5', R = 7-MeO). Addnl. 9 I I and their disodium salts, and 9 II are described.

AN 1971:111916 CAPLUS Full-text

DN 74:111916

OREF 74:18129a,18132a

TI Bischromonyloxy compositions for inhibiting the effects of antigen antibodies

PA Fisons Pharmaceuticals Ltd.

SO Fr. M., 17 pp.

CODEN: FMXXAJ

DT Patent

LA French

FAN.CNT 1

PATENT NO. KIND DATE APPLICATION NO. DATE

PI FR 6893 19690604 FR GB 19660705 GB 19660714

OS MARPAT 74:111916 II 23915-70-0P 23915-71-1P 23937-54-4P 23937-89-5P 31437-68-0P 31545-45-6P

RL: SPN (Synthetic preparation); PREP (Preparation)

(preparation of)

RN 23915-70-0 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid,

5,5'-[(2-hydroxytrimethylene)dioxy]bis[8-allyl-4-oxo-, diethyl ester (8CI) (CA INDEX NAME)

PAGE 1-A

PAGE 2-A

RN 23915-71-1 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid,

5,5'-[(2-hydroxytrimethylene)dioxy]bis[8-allyl-4-oxo- (8CI) (CA INDEX NAME)

PAGE 1-A

23937-54-4 CAPLUS RN

CN 4H-1-Benzopyran-2-carboxylic acid,

5,5'-[(2-hydroxytrimethylene)dioxy]bis[7-(2-hydroxypropoxy)-4-oxo- (8CI)

(CA INDEX NAME)

PAGE 2-A

23937-89-5 CAPLUS RN

4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxy-1,3-propanediyl)bis(oxy)]bis[7-methoxy-4-oxo- (CA INDEX CN NAME)

RN 31437-68-0 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxy-1,3-propanediyl)bis(oxy)]bis[7-methoxy-4-oxo-, disodium salt (9C1) (CA INDEX NAME)

RN 31545-45-6 CAPLUS CN 4H-1-Benzopyran-2-

4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxytrimethylene)dioxy]bis[7-(2-hydroxypropoxy)-4-oxo-, disodium salt (8C1) (CA INDEX NAME)

2 Na

ANSWER 2441 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN L4

The action of disodium cromoglycate (I) on reagin (type II) reactions was AB studied in the rat. Passive cutaneous anaphylaxis was substantially inhibited when 0.5 mg I/kg was given i.v. with antigen to rats, and 2 mg gave complete inhibition. To determine the specificity of the drug, rats were sensitized at different times with both rat reagin and rabbit hyperimmune serum. When given with the antigens, 8 mg I/kg completely inhibited both passive cutaneous anaphylaxis and mast cell disruption induced by rat reagin. Smaller doses inhibited only mast cell disruption. The effects of the drug on the release of spasmogens from sensitized rat lung challenged with antigen were studied in the presence of isolated guinea pig ileum. Given with the antigen, 10 µg I/ml caused an average of 40% reduction in the height of ileal contraction, while 100 ug/ml caused a 64% reduction. The drug had no effect on the response to spasmogens added to the bath; thus, it must have inhibited the antigen-induced release of these substances. The results show that I possesses unusual activity in inhibiting reagin reactions and should be useful both as a drug and in the study of allergies.

AN 1971:97144 CAPLUS Full-text

DN 74:97144

OREF 74:15791a,15794a

- Disodium cromoglycate a specific inhibitor of certain reagin (type II) antibody-antigen reactions
- AU Blair, A. M. J. N.; Clarke, Alan James
- CS Fisons Pharm. Res. Lab., Loughborough/Leicester, UK
- Cell. Humoral Mech. Anaphylaxis Allergy, Proc. Int. Symp. Can. Soc. SO Immunol., 3rd (1969), Meeting Date 1968, 114-18. Editor(s): Movat, H. Z. Publisher: Karger, Basel, Switz. CODEN: 22XMAR
- Conference DT T.A English
- 15826-37-6

RL: BIOL (Biological study) (reaginic antibody-antigen reaction in relation to)

- RN 15826-37-6 CAPLUS
- 4H-1-Benzopyran-2-carboxylic acid, CN

5,5'-[(2-hvdroxv-1,3-propanediv1)bis(oxv)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

- L4 ANSWER 2442 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN
- GI For diagram(s), see printed CA Issue.
- AB In vitro studies using rat s.c. connective tissue sensitized with rat reagin revealed that disodium cromoglycate (I) inhibited the allergic release of histamine if present during antigenic challenge, but when present during sensitization had no effect on antigen-induced release of histamine provided that I was removed prior to challenge. Tissue which had undergone a primary antigen challenge in the presence of I failed to release histamine upon removal of I and rechallenge, indicatng that antigen/antibody interaction occurred in the presence of the compound, resulting in desensitization to a subsequent antigen challenge. Results from in vivo passive cutaneous anaphylactic(PCA) reactions using tissue sites sensitized with 2 reaginic antibodies, which permitted a sequence of antigen challenges, demonstrated that it was possible to desensitize tissue, without the release of the anaphylaxis mediators, by an antigen challenge and I treatment. In these sites sensitized by 2 antibodies, the immunol. reactivity was maintained following a primary antigen challenge and I treatment, as a subsequent challenge with the dissimilar antigen produced a good PCA reaction. Thus, I may act directly or indirectly at a stage following antigen antibody reaction, but prior to the release of anaphylaxis mediators.
- AN 1971:40889 CAPLUS Full-text
- DN 74:40889
- OREF 74:6585a,6588a
- TI Mode of action of disodium cromoglycate. Studies on immediate type hypersensitivity reactions using 'double sensitization' with two antigenically distinct rat reacins.
- AU Orr, Thomas S. C.; Pollard, M. C.; Gwilliam, Jessie; Cox, James S. G.
- CS Res. Dev. Lab., Fisons Pharm. Ltd., Loughborough/Leicestershire, UK
- SO Clinical and Experimental Immunology (1970), 7(5), 745-57 CODEN: CEXIAL: ISSN: 0009-9104
- DT Journal
- LA English
 - T 15826-37-6
 - RL: BIOL (Biological study)
 - (histamine release response to, in allergy)
- RN 15826-37-6 CAPLUS
- CN 4H-1-Benzopyran-2-carboxylic acid,
 - 5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

osc.G THERE ARE 4 CAPLUS RECORDS THAT CITE THIS RECORD (4 CITINGS)

- L4 ANSWER 2443 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN
- AB Release of anaphylactic mediator (AM) in vitro from lung slices of egg albumen-sensitized quinea pig after addition of egg albumen was not affected by the presence of cromoglycate (I). Incubation of I with the liver homogenates of the untreated quinea pig could not convert I to any active derivative stimulating the AM release. A slight inhibition of AM release by I was found when the quinea pig was sensitized by egg albumen supplemented with Freund's adjuvant. When monkeys (Macaca irus) had been sensitized by the serum of a human with housedust allergy, the AM release from the lung slices upon adding the reagin-containing serum was significantly inhibited by I. The inhibitory action of I is associated with the reagin-anaphylactic system.
- AN 1971:21708 CAPLUS Full-text
- DN 74:21708
- OREF 74:3499a,3502a
- Effect of disodium cromoglycate on the anaphylactic mediator release from lung of some animal species
- Koda, Akihide; Nagai, Hiroichi; Hiramatsu, Masahiko; Katsura, Eiji AU
- CS Gifu Pharm. Coll., Gifu City, Japan
- SO Arerugi (1970), 19(8), 597-604
- CODEN: ARERAM; ISSN: 0021-4884
- DT Journal
- LA Japanese
- IT 15826-37-6
 - RL: BIOL (Biological study) (anaphylaxis response to)
- RN 15826-37-6 CAPLUS
- 4H-1-Benzopyran-2-carboxylic acid,
 - 5.5'-[(2-hvdroxv-1.3-propanediv1)bis(oxv)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

L4 ANSWER 2444 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN

AB The 4 hr passive cutaneous anaphylaxis reaction (PCA) in rats induced by rat anti-dinitrophenyl 78y2 antibody has been investigated and compared with the rat reagin-induced PCA reaction. Time course studies revealed that the PCA reaction was made up of at least 2 parts, an early immediate reaction which involved mast cell degranulation, was inhibited by disodium cromoglycate and by cyproheptadine and a late reaction unaffected by disodium cromoglycate, by cyproheptadine, and by an anti-SR5-A agent, diethylcarbamazine. The early part of the 78 $\gamma 2$ reaction was comparable to the rat reagin PCA reaction whereas the later part of the 78 $\gamma 2$ reaction does not appear to involve similar pathways or mediators.

AN 1970:496795 CAPLUS Full-text

DN 73:96795

OREF 73:15799a,15802a

TI Passive cutaneous anaphylaxis in the rat with disodium cromoglycate. I. Cutaneous reactions induced by an anti-DNP 7Syz antibody

AU Orr, Thomas S. C.; Gwilliam, Jessie; Cox, James S. G.

CS Res. Develop. Lab., Fisons Pharm. Ltd., Loughborough, UK

SO Immunology (1970), 19(3), 469-79

CODEN: IMMUAM; ISSN: 0019-2805

DT Journal

LA English

IT 15826-37-6

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(immunosuppressant activity of, anaphylaxis in relation to)

RN 15826-37-6 CAPLUS CN 4H-1-Benzopyran-2-

4H-1-Benzopyran-2-carboxylic acid,

5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

osc.g 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)

ANSWER 2445 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN T. 4

AB Intracardiac injection of disodium cromoglycate (I) into the rat (2-8 mg/kg) inhibited passive cutaneous anaphylaxis and mast cell disruption induced by anti-egg albumin antiserum. Suppression of liberation of the antigen-antibody reaction mediator by I is suggested.

AN 1970:453970 CAPLUS Full-text

DN 73:53970

OREF 73:8875a,8878a

Inhibiting effect of disodium cromoglycate on passive cutaneous anaphylaxis in rats

AU Sudo, Morio; Yoshida, Toru

CS Iwate Med. Coll., Morioka, Japan

Arerugi (1970), 19(4), 250-9 SO CODEN: ARERAM; ISSN: 0021-4884

DT Journal

LA Japanese

ΙT 15826-37-6

> RL: BIOL (Biological study) (anaphylaxis inhibition by)

RN 15826-37-6 CAPLUS

4H-1-Benzopyran-2-carboxylic acid, CN

5,5'-[(2-hydroxy-1,3-propanediyl)bis(oxy)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

L4 ANSWER 2446 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN

The possibility that disodium cromoglycate (1,3-bis(2-carboxychromon-5-yloxy)-2-hydroxypropane) (I) inhibits bronchoconstriction in humans with extrinsic asthma by specifically suppressing anaphylactic reactions initiated by interaction of antigen with antibodies was studied in rats. Passive cutaneous anaphylaxis was induced in rats by i.v. injecting antigenic worm exts. or dinitrophenylated protein conjugates 48 and 4 hr, resp., after the intradermal administration of rat antiserum containing anti-Nippost rongylus brasiliensis antibodies or an IqGa fraction from rat antiserum containing antidinitrophenyl bovine y globulin. The simultaneous (in sep. syringes) injection of 25 mg I with the worm extract abolished the cutaneous anaphylactic reaction to this antigen; I also abolished the reaction when given immediately prior to the antigen. I partially inhibited the 4-hr passive cutaneous anaphylactic reaction in rats pretreated with IgGa antibodies. Since the 48-hr anaphylactic reaction is presumed to be mediated by the release of histamine or serotonin from sensitized mast cells upon contact with antigen, it is suggested that I inhibits the immunologic release of histamine and serotonin from these cells regardless of the immunoglobulin class of antibodies involved.

AN 1970:53175 CAPLUS Full-text

DN 72:53175

AB

OREF 72:9719a,9722a

- TI Effect of disodium cromoglycate on certain passive cutaneous anaphylactic reactions
- AU Lopez, Manuel; Bloch, Kurt J.
- CS Allergy and Arthritis Units, Massachusetts Gen. Hosp., Boston, MA, USA
- SO Journal of Immunology (1969), 103(6), 1428-30
- CODEN: JOIMA3; ISSN: 0022-1767 DT Journal
- LA English
 - IT 15826-37-6
 - RL: BIOL (Biological study)
 - (anaphylaxis inhibition by, antibody types in relation to)
- RN 15826-37-6 CAPLUS
- CN 4H-1-Benzopyran-2-carboxylic acid,
 - 5,5'-[(2-hydroxy-1,3-propanediyl)bis(oxy)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)

L4 ANSWER 2447 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN

- AB In the rat disodium cromoglycate showed a distinct antagonistic effect against noradrenaline and isopropylnoradrenaline and a weak antagonistic effect against histamine and bradykinin but it did not interfere with the effect of serotonin, with the amine release elicited by compound 1935 L, with the hypotension produced by kinin-forming substances such as ellagic acid or dextran sulfate, or with the development and course of anaphylactic shock.
- AN 1970:41343 CAPLUS Full-text
- DN 72:41343
- OREF 72:7575a,7578a
- TI Pharmacological properties of disodium cromoglycate [disodium 1,3-bis(2-carboxychromon-5-yloxy)-2-hydroxypropane tetrahydrate] in the rat
- AU Lecomte, Jean
- CS Univ. Liege, Liege, Belg.
- SO Acta Allergologica (1969), 24(3), 226-31
- CODEN: ACALAF; ISSN: 0001-5148
- DT Journal
- LA French
 - I 15826-37-6

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

- (pharmacology of)
- RN 15826-37-6 CAPLUS
- CN 4H-1-Benzopyran-2-carboxylic acid,
 - 5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

L4 ANSWER 2448 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN

AB Three in vitro models of the immediate hypersensitivity reaction, two involving reaginic antibodies in human lung and one involving reaginilke antibodies in rat lung, were used in laboratory investigations on disodium cromoglycate. The release of spasmogens after antibody-antigen reaction in each of these models was partially inhibited by disodium cromoglycate and it is suggested that this effect may be related to the clin. efficacy of the compound in allergic asthma.

AN 1970:29791 CAPLUS Full-text

DN 72:29791

OREF 72:5421a,5424a

TI Disodium cromoglycate. Activity in three in vitro models of the immediate hypersensitivity reaction in lung

AU Sheard, Philip; Blair, A. M. J. N.

S Fisons Pharm. Res. Lab., Loughborough, UK

SO International Archives of Allergy and Applied Immunology (1970), 38(2), 217-24 CODEN: IAAAAM, ISSN: 0020-5915

DT Journal

LA English

15826-37-6

RL: BIOL (Biological study)

(hypersensitivity inhibition by, in lungs in allergy)

RN 15826-37-6 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid,

5,5'-[(2-hydroxy-1,3-propanediyl)bis(oxy)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

OSC.G 8 THERE ARE 8 CAPLUS RECORDS THAT CITE THIS RECORD (8 CITINGS)

L4 ANSWER 2449 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN

- AB Cromoglycate has been shown to be a potent inhibitor of histamine release by antigen from passively sensitized lung of man and monkey. Slight inhibition of histamine release was also produced in lung of actively sensitized monkey and in actively sensitized leukocytes of man. Reactions not consistently inhibited were: direct and Prausnitz-Kuestner skin tests in man, and histamine release from actively sensitized leukocytes of rabbit. There was no inhibition of the anaphylactic histamine release from guinea-pig lung, whether actively or passively sensitized.
- AN 1970:11025 CAPLUS Full-text
- DN 72:11025
- OREF 72:1987a,1990a
- TI Inhibition of allergic reactions in man and other species by cromoglycate
- AU Assem, E. S. K.; Mongar, J. L.
- CS Dep. Pharmacol., Univ. Coll. London, London, UK
- SO International Archives of Allergy and Applied Immunology (1970), 38(1), 68-77
 - CODEN: IAAAAM; ISSN: 0020-5915 T Journal
- DT Journal LA English
- IT 15826-37-6
 - RL: BIOL (Biological study)
 - (histamine liberation inhibition by, in allergy)
- RN 15826-37-6 CAPLUS
- CN 4H-1-Benzopyran-2-carboxylic acid,
 - 5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

L4 ANSWER 2450 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN

GI For diagram(s), see printed CA Issue.

Compds. I, II, and III are prepd. from IV compds., where one of R2 and R3 is OH and one of R1, R3, and R4 is Ac. Thus, a mixture of 4,2,6-MeO-(HO)2C6H2Ac 9.1, epichlorohydrin 2.33 parts, NaOEt, and EtOH is refluxed 4.5 hrs. to give 5.45 parts 1,3-bis(2-acetyl-3-hydroxy-5-methoxyphenoxy)-2-propanol (V), m. 180-2°. A mixture of V 2.1, EtO2CCO2Et 3.7, and dioxane 30 is added to a solution of NaOEt in EtoH (prepared from Na 0.92 and EtoH 20), EtoH 20 parts is added, and the mixture is refluxed 4 hrs. and worked up to give 1.73 parts 1,3-bis(2-carboxy-7-methoxy-5-chromonyloxy)-2-propanol (VI), m. 245° (decomposition). Similarly prepared are the following I (R1 = H) (R, R2, R3, and m.p. given): (CH2)3, OMe, H, 288° (monohydrate); CHOH, OCH2CH(OH)Me, H, $244-6^{\circ}$; CHOH, H, allyl, $214-18^{\circ}$ (decomposition); the following II (R1 = H) (R, R2, R3, and m.p. given): CHOH, OCH2CH(OH)Me, H, 160-1° (decomposition); (CH2)3, OMe, H, 159-70°; CHOH, H, allyl, 210-30°; the following III (R = H) (R1 and m.p. given): Me, 258-60° (dihydrate); CH2CH(OH)Me, 180-3°; and the following IV (R, R1, R2, R3, R4, and m.p. given): (CH2)3, Ac, OH, H, OMe, 146-7°; CHOH, Ac, OH, Ac, OH, 244-5°; CHOH, H, OH, Ac, OH, 251-3°; CHOH, H, OH, Ac, OCH2CH(OH)Me, 148-50°; CHOH, Ac, OH, H, OCH2CH(OH)Me, 201-3°; (CH2)3, H, OH, Ac, OMe, 130-2°; (CH2)3, OMe, H, OH, Ac, 145-8°; (CH2)3, OCH2CH(OH)Me, H, OH, Ac, 122°; CHOH, Ac, OH, allyl, H, -; CHOH, allyl, OH, Ac, H, 137-9°. II (R = CHOH, R1 = R3 = H, R2 = OCH2Ph) is heated with HBr in HOAc to give II (R = CHOH, R1 = R3 = H, R2 = OH); monohydrate m. 245-6°. VI and the I (R1 = H), II (R1 = H), and III (R = H) prepared are converted to the corresponding I (R1 = Na), II (R1 = Na), and III (R = Na). I (R = CHOH, R1 = Et, R2 = H, R2 = ally1) (m. 153-5°) is obtained by the EtO2CCO2Et cyclization reaction and converted to the I (R1 = H) compound Also prepared, according to known methods, are the following intermediates (m.p. given): 2,4-diacety1-5-(2hydroxypropoxy)resorcinol, 152-4°; 2,6,4-(HO)2[MeCH(OH)CH2O]C6H2Ac, 177-8°; 2,5,4-[HO)2[MeCH(OH)CH2O]C6H2Ac, 186-8°. III [R = Et, R1 = CH2CH(OH)Me] (m. 180-3°) is also obtained and converted to the III (R = H) compound 5.7-Dihydroxy-2-methylchromone is treated with Br(CH2)5Br to give 1,5-bis(5hydroxy-2-methyl-7-chromonyloxy)pentane (m. 185°) which is converted to the di-Me ether (m. 208-9°); the ether is heated with KOH in EtOH to give IV [R = (CH2)3, R1 = H, R2 = OH, R3 = Ac, R4 = OMe], m. $130-2^{\circ}$.

AN 1969:501709 CAPLUS Full-text

DN 71:101709

AB

OREF 71:18940h,18941a

TI α, ω-Bis(2-carboxychromonyloxy) alkanes

PA Fisons Pharmaceuticals Ltd.

SO Fr., 17 pp.

CODEN: FRXXAK

DT Patent LA French

FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. DATE PΙ FR 1533506 19680719 FR 1967-112960 19670704 GB 19660705 GB 19660714 DE 1593882 DE GB 1190193 GB GB 1190194 GB US 3519652 19700707 19670703 HS US 3705945 19721212 US 19700507

23915-70-0P 23915-71-1P 23937-54-4P IT

23937-89-5P

RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of)

23915-70-0 CAPLUS RN

CN 4H-1-Benzopyran-2-carboxylic acid,

5,5'-[(2-hydroxytrimethylene)dioxy]bis[8-allyl-4-oxo-, diethyl ester (8CI) (CA INDEX NAME)

PAGE 2-A

RN 23915-71-1 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid,

> 5,5'-[(2-hydroxytrimethylene)dioxy]bis[8-allyl-4-oxo- (8CI) (CA INDEX NAME)

PAGE 2-A

RN 23937-54-4 CAPLUS

2N 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxytrimethylene)dioxy]bis[7-(2-hydroxypropoxy)-4-oxo-(8CI) (CA INDEX NAME)

PAGE 1-A

PAGE 2-A

5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[7-methoxy-4-oxo- (CA INDEX NAME)

L4 ANSWER 2451 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN

GI For diagram(s), see printed CA Issue.

AB Diethers ArOXAr (I) where Ar is a 2-acety1-3-hydroxyphenyl group, are treated with Eto2CcO2Et to give II; III, IV, and V are also prepared, V are prepared from ArOXOAr1 (VI). Thus, a solution of 1,3-bis(2-acety1-3-hydroxyphenoxy)propane (VII) 6.9 in Eto2CCO2Et 15 is added to a solution of Na 3 in EtoH 30 and C6H6 50 parts; the mixture is refluxed 20 hrs. to give 4.5 parts 1,3-bis(2-carboxychromon-5-yloxy)propane (VIII) di-Et ester, m. 182-3° which

R1, R2, R3, m.p. or decomposition pt.* given): CH2CH(OH)CH2, CO2Et, H, H, 180-2°; CH2CH(OH)CH2, CO2H, H, H, 241-2°*; CH2CH:CHCH2, CO2Et, H, H, 216-17°; CH2CH:CHCH2, CO2Et, H, H, 193-5° (monohydrate); CH2CH(OH)CH2O(CH2)*40CH2CH(OH)CH2, CO2H, H, H, 80°* (dihydrate); (CH2)4, CO2Et, H, H, 19-6°); (CH2)4, CO2H, H, H, 288-30° (monohydrate); (CH2)5, CO2Et, H, H, 150-2°; (CH2)5, CO2Et, H, H, 152-8°; (CH2)6, CO2H, H, H, 288-30° (monohydrate); (CH2)6, CO2Et, H, H, 154.5°; (CH2)6, CO2H, H, H, 288-30° (monohydrate); CH2)6, CO2Et, H, H, 164.5-8°; (CH2)10, CO2Na, H, H, -7 CH2CH(OH)CH2CCH2CH(OH)CH2, CO2H, H, H, 216-18° (monohydrate); CH2CH2OCH2CH2, CO2Et, H, H, 199-31.5°; CH2CH2OCH2CH2, CO2H, H, 219-20°; CH2CH(OH)CH(OH)CH(OH)CH2, CO2Et, H, H, 216-17°; CH2CH(OH)CH2CH2, CO2Et, H, H, 216-17°; CH2CH(OH)CH2CH2, CO2H, H, H, 260-2° (dihydrate); CH2CH(OH)CH2CH2, CO2Et, H, H, 216-17°; CH2CH(OH)CH2CH2, CO2H, H, H, 260-2° (dihydrate); CH2CH(OH)CH2CH2, CO2Et, H, H, 161-17°; CH2CH(OH)CH2CH2, CO2H, H, L264-7° (monohydrate); (CH2)5, CO2Et, H, C1, 162-4°; (CH2)5, CO2H, H, C1, 244°; CH2CH(OH)CH2, CO2Et, M, H, 194-6°;

is converted to VIII di-Na salt. Similarly prepared are the following II (X,

H, 219-20°, CH2CH(OH)CH2(OH)CH2. CO2EE, H, H, 224-6°, CH2CH(OH)CH2(OH)CH2(OH)CH2, CO2H, H, H, 260-2° (dihydrate); CH2CH(OH)CH2CH2, CO2EE, H, H, 216-17°, CH2CH(OH)CH2CH2, CO2H, H, H, 226-7° (monohydrate); CH2D5, CO2EE, H, C1, 162-4°; (CH2D5, CO2H, H, C1, 244°; CH2CH(OH)CH2, CO2EE, Me, H, 194-6°; CH2CH(OH)CH2, CO2EE, Me, H, 240-1° (monohydrate); CH2CH(OH)CH2, CO2EE, H, Et, 159-61°; CH2CH(OH)CH2, CO2H, H, BE, 193-4° (dihydrate); CH2CH2CHMECH2CH2, CO2EE, H, H, 128-30°, CH2CH2CHMECH2CH2, CO2H, H, B, 128-17° (monohydrate); ophenylene, CO2EE, H, H, 128-30°, CH2CH2CHMECH2CH2, CO2H, H, B, 125-17°, Gmonohydrate); ophenylene, CO2EE, H, H, 128-30°, CH2CH2CHMECH2CH2, CO2H, H, H, (tetrahydrate); ophenylene, CO2EE, H, H, H, 183-7°; CH2CH2, CO2E, H, H, 123-7°; CH2CH2, CO2E, H, H, 128-7°; CH2CH2, CO2E, H, H, 168-8°; CH2C(H2CH2) (CH2CH)CH2C, CO2EC, H, H, 168-8°; CH2C(H2CH2) (CH2CH)CH2C, CO2EC, H, H, -285°; CH2C), Me, H, H, 140-3°; CH2CH3, H, H, Me, 274-6°; CH2CH(OH)CH2, Et, H, Me, 272-5° (dihydrate); CH2CH3, Et, H, Me, 191-3°; CH2CH(OH)CH2, Et, H, Me, 272-5° (dihydrate); CH2C), Et, -1, (CH2)5, H, H, 25-7°; CH2CH(OH)CH2, Et, C), CH2CH, CHCHOH)CH2, Et, C), CH2CH, CHCHOH)CH2, Et, C)

187-9°; CH2CH(OH)CH2, H, 268-70° (dihydrate); the following V (X, R1, R2, and m.p. given): CH2CH(OH)CH2, 2-eth-oxycarbonylchromon-7-yl, Et, 193-4.5°; CH2CH(OH)CH2, 2-carboxychromon-7-yl, H, 194-200°; (CH2)5, 2-ethoxycarbonylchromon-7-yl, Et, 149-52°; (CH2)5, 2-carboxychromon-7-yl, H, 249-51°; CH2CH(OH)CH2, 2-ethoxycarbonyl-8-ethylchromon-5-yl, Et, 166-6.5°; CH2CH(OH)CH2, 2-carboxy-8-ethylchromon-5-yl, H, 190-2° (dihydrate); CH2CH(OH)CH2, 2-ethoxycarbony1-6-chlorochromon-7-y1, Et, 166-8°; CH2CH(OH)CH2, 2-carboxy-6-chlorochromon-7-yl (Na salt), Na, -; CH2CH(OH)CH2, 2ethoxycarbonylchromon-6-vl, Et. 164-6°: CH2CH(OH)CH2, 2-ethoxycarbonylchromon-8-yl, Et, 166-9°; 1,5-bis(2-carboxychromon-8-yloxy)pentane (IX) di-Et ester (m. 128-30°), IX monohydrate (m. 237-8°), and the di-Na salt, Ca salt, Mg salt, and dipiperidine salt of II [X = CH2CH(OH)CH2, R1 = CO2H, R2 = R3 = H]. II [X = CH2CH(OH)CH2OCH(CH2OH)CH2, R1 = CO2H, R2 = R3 = H] (m. 276-80°) is obtained as by-product in the preparation of the II (X = dioxan-2,5dividimethylene) compound II [X = (CH2)5, R1 = Me, R2 = R3 = H] (X) 5 is heated with SeO2 6 in dioxane 100 parts to give II [X = (CH2)5, R1 = CO2H, R2 = R3 = H1 monohydrate (XI), m. 226-8°. X 2.7 is treated with BzH 1.5 and Na 0.294 parts in EtOH to give II [X = (CH2)5, R1 = CH:CHPh, R2 = R3 = H] (m. 217-20°) which is oxidized (KMnO4) to give XI, m. 226-8°. 1,4-Bis(4hydroxyphenoxy)pentane (XII) 5.8 is treated with EtO2CC.tplbond.CCO2Et 6.8 parts to give 1.5 parts IV [X = (CH2)5, R1 = H] m. 270-1°. Also prepared, according to known methods, are the intermediates, VII (m. 184-5°) and XII (m. 110-12°), the following I intermediates (X, Ar, and m.p. given): CH2CH(OH)CH2, 2,3-Ac(HO)C6H3 (Q), 165-6°; CH2CH:CHCH2, Q, 145-6°; CH2CH(OH)CH2O(CH2)4OCH2CH(OH)CH2, Q, -; (CH2)4, Q, 219-21°; (CH2)5, Q, 131-3°; (CH2)6, Q, 147.5-8.5°; (CH2)10, Q, 102.5-4°; CH2CH(OH)CH2OCH2CH(OH)CH2, Q, 129-31°; CH2CH2OCH2CH2, Q, 120.5-1.5°; CH2CH(OH)CH(OH)CH2, Q, 211-12°; CH2CH(OH)CH2CH2, O, 207.5-8.5°; (CH2)5, 4,2,3-C1(Ac)(H0)C6H2, 96°; CH2CH(OH)CH2, 5,2,3-Me(Ac)(HO)C6H2, 185-6°; CH2CH(OH)CH2, 4,2,3-Et(Ac)(HO)C6H2, 135-7°; (CH2)5, 3,2-Ac(HO)C6H3, 103.5-4.5°; (CH2)5, 2,4,3-Me(Ac)(HO)C6H2(R), 116-17°; CH2CH(OH)CH2, R, 151-3°; CH2CH2CHMeCH2CH2, Q, 123-4°; CH2CH(OH)CH2, 2,4,5-Cl(Ac)(HO)C6H2, 197-9°; (CH2)5, 3,4-Ac(HO)C6H3 (S), 107-9°; o-phenylene, Q, 148-53°; CH2CH(OH)CH2, S, 127-9°; (CH2)8, Q, 107-9°; (CH2)9, Q, 55-9°; CH2CH2, Q, 188-9°; oxetan-3,3-diyldimethylene, Q, 209-11°; dioxan-2,5-diyldimethylene, Q, 230-2°; the following VI (X = CH2CH(OH)CH2] (Ar, Ar1, and m.p. given): Q, 4,3-Ac(HO)C6H3, 182-5°; Q, 4,2,3-Et(Ac)(HO)C6H2, 102-3°; Q, 2,4,5-C1(Ac)(HO)C6H2, 139-40°; Q, S, 184-5°; Q, 3,2-Ac-(HO)C6H3, 166-9°; and the following intermediates (m.p. given): VI [X = (CH2) 5, Ar = 2,3-Ac(HO)C6H3, Ar1 = 4,3-Ac(HO)C6H3] 91-1.5°; 2-(2,3epoxypropoxy)-6-hydroxyacetophenone, 61-3°; 6,2-HO[C1CH2CH(OH)CH2O1C6H3Ac. b1.5 166-8°; 5-(2,3-epoxypropoxy)-2-hydroxyacetophenone, 76.9°.

AN 1969:491309 CAPLUS Full-text

DN 71:91309

OREF 71:16994h,16995a

TΙ α , ω -Bis(2-carboxychromon-5-yloxy) alkanes

TN Fitzmaurice, Colin; Lee, Thomas Brian

PA Fisons Pharmaceutical Ltd.

SO Brit., 31 pp. Division of Brit. 1144905 CODEN: BRXXAA

DT Patent

LA English

FAN CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE	
PI	GB 1144906		19690312	GB 1968-37765	19650325	
	DE 1918142			DE		
	DE 1920365			DE		
OS	MARPAT 71:91309					

IT 15826-37-6P 16110-51-3P 16119-86-5P 16129-38-7P 16130-23-7P 16130-25-9P 16139-24-5P 16139-25-6P 16146-53-5P

16150-45-1P 23874-48-3P

RL: SPN (Synthetic preparation); PREP (Preparation)

(preparation of)

15826-37-6 CAPLUS RN

CN 4H-1-Benzopyran-2-carboxylic acid,

5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

RN 16110-51-3 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid,

5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[4-oxo- (CA INDEX NAME)

RN 16129-86-5 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxytrimethylene)dioxy]bis[4-oxo-, calcium salt (1:1) (8CI) (CA INDEX NAME)

RN 16129-88-7 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxy-1,3-propanediyl)bis(oxy)]bis[4-oxo-, compd. with piperidine (1:2) (9C1) (CA INDEX NAME)

CM 1

CRN 16110-51-3 CMF C23 H16 O11

CM 2

CRN 110-89-4 CMF C5 H11 N RN 16130-23-7 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[7-methyl-4-oxo- (CA INDEX NAME)

RN 16130-25-9 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxytrimethylene)dioxy]bis[8-ethyl-4-oxo-, diethyl ester (8CI) (CA INDEX NAME)

RN 16139-24-5 CAPLUS

CN

4H-1-Benzopyran-2-carboxylic acid, 5-[3-[[2-(ethoxycarbonyl)-4-oxo-4H-1-benzopyran-5-yl]oxy]-2hydroxypropoxy]-8-ethyl-4-oxo-, ethyl ester (CA INDEX NAME)

- RN 16139-25-6 CAPLUS
- CN 4H-1-Benzopyran-2-carboxylic acid, 5-[3-[(2-carboxy-4-oxo-4H-1-benzopyran-5-y1)oxy]-2-hydroxypropoxy]-8-ethyl-4-oxo (CA INDEX NAME)

- RN 16146-53-5 CAPLUS
- CN 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[8-ethyl-4-oxo- (CA INDEX NAME)

RN 16150-45-1 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxy-1,3-propanediyl)bis(oxy)]bis[4-oxo-, 2,2'-diethyl ester (CA INDEX NAME)

RN 23874-48-8 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxytrimethylene)dioxy]bis[7-methyl-4-oxo-, diethyl ester (8CI) (CA INDEX NAME)

$$\begin{array}{c} \text{Me} & \begin{array}{c} 0 \\ \text{OEt} \\ \end{array} \\ \text{OCH}_2 \\ \text{CH}_2 \\ \text{CH}_2 \\ \text{OH} \end{array}$$

OSC.G 4 THERE ARE 4 CAPLUS RECORDS THAT CITE THIS RECORD (4 CITINGS)

- L4 ANSWER 2452 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN
- GI For diagram(s), see printed CA Issue.

The title compds. (I) were prepd. by condensing two mols. II with a compd. AXB AB in a 1 or 2 step reaction. Thus, 4-methyl-7-hydroxy-8-acetylcoumarin 109, K2CO3 35, epichlorohydrin 26, iso-PrOH 1250, and 40% aqueous PhCH2NMe3OH 1 was refluxed 65 hrs. with stirring, iso-PrOH 1000 distilled, and the residue diluted with H2O 500 to give I [R = Me, R1 = Ac, R2 = R3 = H, X = CH2CH(OH)CH2] 16 parts by weight, m. 234° (EtOH). Similarly prepared were I (R, R1, R2, R3, X, m.p., and % yield given): Me, H, H, AcO, (CH2)5, 205-8° (C6H6-CHCl3), -; Me, H, H, H, (CH2)5, 176-8° (MeOCH2CH2OH), 95; Me, H, H, H, CH2CH(OH)CH2, 110-2° (MeOCH2CH2OH), 35. Also prepared were 1.3-bis(2-acetyl-3-hydroxyphenoxy)-2-propanol, m. 160-5° (EtOH), 1,3-bis(2-carboxychromon-5yloxy)-2-propanol di-Et ester, m. 180-2° (EtOH-C6H6), and the di-Na salt of 1.3-bis(2-carboxychromon-5-yloxy)-2-propanol monohydrate. AN

1969:481181 CAPLUS Full-text

DN 71:81181

OREF 71:15037a,15040a

TΙ 7,7'-(Polymethylenedioxy)bis(4-methylcoumarins)

IN Fitzmaurice, Colin; Lee, Thomas Brian

PA Fisons Pharmaceuticals Ltd.

so Ger. Offen., 16 pp.

CODEN: GWXXBX DT Patent

LA German

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	DE 1810064		19690619	DE 1968-1810064	19681120
				GB	19671122
	FR 1598304			FR	
	GB 1237878			GB	
	US 3567741		19710302	US	19681121
	ZA 6807363		19680000	ZA	
- m	10000 00 00 1010	2 25 25			

15826-37-6P 16150-45-1P

RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of)

RN 15826-37-6 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid,

5,5'-[(2-hydroxy-1,3-propanediyl)bis(oxy)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[4-oxo-, 2,2'-diethyl ester (CA INDEX NAME)

L4 ANSWER 2453 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN

Phospholipase A (10 µg./ml.) degranulated 62.5% of the mast cells from rat AB connective tissue in 15 min. at 37.5°; in the presence of 10 µg. disodium cromoglycate (I)/ml., however, only 42.6% of the mast cells were degranulated by phospholipase A. In the reagin antibody/antigen system, 55.4% of the mast cells were degranulated; this was reduced to 32.6% in the presence of I. Addition of I also reduced the histamine release from mast cells by phospholipase A from 21.5 to 7.3%; in the reaginic antibody/antigen reaction, I reduced the release from 31 to 14.9%. I may act on a phospholipid enzyme or some other antibody/antigen activated enzyme or enzymes and thus inhibit mast cell degranulation.

AN 1969:447988 CAPLUS Full-text

DN 71:47988

OREF 71:8811a,8814a

Disodium cromoglycate, an inhibitor of mast cell degranulation and histamine release induced by phospholipase A

ΑU Orr, Thomas S. C.; Cox, J. S. G.

CS Res. Develop. Lab., Fisons Pharm. Ltd., Loughborough, UK

SO Nature (London, United Kingdom) (1969), 223(5202), 197-8

CODEN: NATUAS: ISSN: 0028-0836

Journal English LA

DT

15826-37-6

RL: BIOL (Biological study)

(mast cells degranulation blocking by)

RN 15826-37-6 CAPLUS

4H-1-Benzopyran-2-carboxylic acid, CN

5,5'-[(2-hydroxy-1,3-propanediyl)bis(oxy)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

OSC.G 10 THERE ARE 10 CAPLUS RECORDS THAT CITE THIS RECORD (10 CITINGS)

L4 ANSWER 2454 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN

AB Studies show max. histamine release at 37-40°. At upper and lower limits release ranges from 4-11%. Ca ions are required to obtain release. Absence of Mg from the reaction medium does not inhibit histamine release. Disodium cromoglycate acts as an inhibitor and also behaves as a weak histamine liberator.

AN 1969:447973 CAPLUS Full-text

DN 71:47973

OREF 71:8807a,8810a

TI Effect of disodium cromoglycate and other inhibitors on in vitro anaphylactic histamine release from guinea pig basophil leukocytes AU Greaves, M. W.

CS Roy. Victoria Infirmary, Newcastle upon Tyne, UK

SO International Archives of Allergy and Applied Immunology (1969), 36, 497-505

CODEN: IAAAAM; ISSN: 0020-5915

DT Journal

LA English

IT 15826-37-6

RL: BIOL (Biological study)

(histamine in basophils after treatment with, in anaphylaxis)

RN 15826-37-6 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid,

5,5'-[(2-hydroxy-1,3-propanediyl)bis(oxy)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

- L4 ANSWER 2455 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN
- AB A comparison between the anaphylaxis induced in rats by rat reagin sera and the cutaneous reaction induced by hyperimmune rabbit anti-ovalbumin serum indicated that the 2 skin responses, of type I and type III, resp., are produced in different ways. The reagin-induced reactions were associated with degranulation of mast cells and were inhibited by disodium cromoglycate, a compound used in treatment of asthma in humans. The disodium cromoglycate interfered with mast cell permeability, apparently after the union of antigen and reagin. It is possible that the critical pathway involved is common to both human and rat reagin systems since both are inhibited by the compound recardless of antien used.
 - 1969:411317 CAPLUS Full-text
- DN 71:11317

AN

- OREF 71:2063a,2066a
 - TI Passive cutaneous anaphylaxis in the rat, induced with two homologous reaginlike antibody sera, and its specific inhibition with disodium cromoglycate
- AU Goose, J.; Blair, A. M. J. N.
- CS Res. Lab., Fisons Pharm., Loughborough, UK
- SO Immunology (1969), 16(6), 749-60
- CODEN: IMMUAM; ISSN: 0019-2805
- DT Journal LA English
- IT 15826-37-6
 - RL: BIOL (Biological study)
 - (as inhibitor of anaphylaxis)
- RN 15826-37-6 CAPLUS
- CN 4H-1-Benzopyran-2-carboxylic acid,
 - 5,5'-[(2-hydroxy-1,3-propanediyl)bis(oxy)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

THERE ARE 8 CAPLUS RECORDS THAT CITE THIS RECORD (8 CITINGS) OSC.G

ANSWER 2456 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN T. 4

AB Diethylcarbamazine, pipecolamide, nicotinamide, isonicotinic acid hydrazide, and N-amidinobenzamide at doses of 20 mg./kg. i.v. 30 sec. before challenge inhibited by 66, 82, 31, 67, and 44%, resp., the antigen-induced release of slow reacting substance of anaphylaxis in the rat. These effective substances at 10mM concns. had no effect on the viability of rat polymorphonuclear leukocytes suspended in Tyrode solution None of these effective compds. at doses of 30 mg./kg. i.p. 30 min. before challenge significantly inhibited the homocytotropic antibody-mediated release of histamine in the rat. Disodium cromoglycate (50 mg./kg. i.v. 30 min. before challenge) suppressed the homocytotropic antibody-mediated release of histamine without inhibiting the antigen-induced release of the slow reacting substance of anaphylaxis in the rat. Neither diethylcarbamazine nor disodium cromoglycate antagonized the pharmacol. activity of histamine or the slow reacting substance of anaphylaxis in the rat. Thus, the antigen-induced release of slow reacting substance of anaphylaxis and the homocytotropic antibody-mediated release of histamine in the rat can be selectively blocked in vivo by pharmacol, agents which act after antigen and antibody interaction but prior to the formation and release of the mediators.

1969:56183 CAPLUS Full-text AN

70:56183 DN

OREF 70:10545a,10548a

ΤI Pharmacologic dissociation of immunologic release of histamine and

slow-reacting substance of anaphylaxis in rats

ΑU Orange, Robert P.; Austen, K. Frank

CS Robert B. Brigham Hosp., Boston, MA, USA

SO Proceedings of the Society for Experimental Biology and Medicine (1968), 129(3), 836-41

CODEN: PSEBAA; ISSN: 0037-9727 Journal

LA English

DT

15826-37-6

RL: BIOL (Biological study)

(in slow-reacting substance release in anaphylaxis)

15826-37-6 CAPLUS RN

4H-1-Benzopyran-2-carboxylic acid,

5,5'-[(2-hvdroxv-1,3-propanediv1)bis(oxv)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

- L4 ANSWER 2457 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN
- AB Inhalation of 20 mg. of the title compd. inhibited all the allergic reactions produced by inhalation of the appropriate allergens in asthmatic patients with various types of allergies. Inhibition of the immediate asthmatic reactions by the drug may be due to interference with the release of histamine and other substances. Inhibition of the late asthmatic and systemic reactions, thought to be precipitin mediated, may in turn result from inhibition of the immediate reaction.
- AN 1968:450916 CAPLUS Full-text
- DN 69:50916
- OREF 69:9499a,9502a
- TI Inhibitory effects of disodium cromoglycate [disodium salt of 1,3-bis(2-carboxy-4-oxochromen-5-yloxy)propan-2-ol] on allergen-inhalation tests
- AU Pepys, J.; Hargreave, F. E.; Chan, Moira; McCarthy, D. S.
- CS Brompton Hosp., London, UK
- SO Lancet (1968), II(7560), 134-7
 - CODEN: LANCAO; ISSN: 0140-6736
- DT Journal LA English
- IT 15826-37-6
 - RL: BIOL (Biological study)
 - (allergen reaction inhibition by)
- RN 15826-37-6 CAPLUS
- CN 4H-1-Benzopyran-2-carboxylic acid,
 - 5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)

L4 ANSWER 2458 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN

AΒ Di-Na cromoglycate (FPL 670) (Intal) (I) inhibited the passive cutaneous anaphylactic (PCA) reactions in monkeys sensitized with human serum containing reagin, when given intradermally with the antigen, but did not affect the skin reactions to intradermal histamine, 5-hydroxytryptamine, or bradykinin. Homologous PCA reactions with reagin-like antibody in rats, using both the egg albumen-Bordetella pertussis and Nippostrongylus brasiliensis systems, were also substantially inhibited by I, although it did not affect the skin lesions induced by compound 48/80. In contrast, in quinea pigs, homologous PCA reactions with precipitating antibody were unaffected, as were aerosol or i.v. antigen-induced bronchospasm, and the release of histamine and SRS-A (slow reacting substance-anaphylaxis) from actively or passively sensitized lung in vitro. When the release of histamine and SRS-A from chopped human lung, passively sensitized with human reaginic serum, was measured after exposure to specific antigen(s) in vitro, I over a narrow concentration range inhibited the release, and 10 μg . of I/ml. reduced by 40% the contractile response in an in vitro system in which contractions of human bronchial chain, exposed to passively sensitized and shocked human lung, were used to simulate the supposed events in an attack of allergic asthma. I had no adverse effect on several in vitro antibody-virus systems including influenza A, polio virus type II, vaccinia, and herpes simplex with human and rabbit antisera. Likewise, no effect was found on the LD50 in mice of mouse-adapted polio virus, nor on their protection by Salk vaccine; I did not interfere with any of the several bacterial applutinating systems tested. I, whose action was distinct from that of corticosteroids, had few general pharmacol. effects, was rapidly excreted, and seemed to have a low order of toxicity. I appears to inhibit specifically the anaphylactic process initiated by interactions of reaginic antibody and antigen. This novel property may permit a more specific treatment of allergic disease, especially of the lung.

AN 1968:38047 CAPLUS Full-text

DN 68:38047

OREF 68:7371a,7374a

TI Disodium cromoglycate (FPL 670) ("Intal"). Specific inhibitor of reaginic antibody-antigen mechanisms

AU Cox, James S. G.

CS Fisons Pharm., Ltd., Holmes Chapel, UK

SO Nature (London, United Kingdom) (1967), 216(5122), 1328-9 CODEN: NATUAS; ISSN: 0028-0836

OT Journal

LA English

AB

IT 15826-37-6

RL: BIOL (Biological study)
(inhibition of antibody-antigen reaction by)

RN 15826-37-6 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid,

5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

OSC.G 44 THERE ARE 44 CAPLUS RECORDS THAT CITE THIS RECORD (44 CITINGS)

L4 ANSWER 2459 OF 2459 CAPLUS COPYRIGHT 2010 ACS on STN

GI For diagram(s), see printed CA Issue.

The title compds. I are inhibitors of certain types of antigen-antibody reactions; they are useful in the treatment of extrinsic allergic asthma or intrinsic asthma or of hay-fever, urticaria, auto-immune diseases or virus infections. They can be compounded with bronchodilators when used in inhalation preparation I are prepared by the reaction of II, III, and XAX' (Y is H, Ac, or a carboxylic ester group; Z is OH; or Y and Z (or Y' and Z') form together a chromonyl-2-carboxylic group, X and X' are reactive derivs. which form the ether bond between the 2 parts of the I mol.). Thus, a mixture of 2,6-(HO)2C6H3Ac 30.4, Br(CH2)3Br 20.2, and powdered K2CO3 12.8 in Me2CO 200 parts is refluxed 72 hrs. The precipitate is filtered and washed with Me2CO and H2O. The combined Me2CO and aqueous washing liquors are evaporated to give an oil, which is boiled with Et2O, the obtained precipitate is combined with the first precipitate and the mixture extracted several days in a Soxhlet apparatus with iso-PrOH to give 16.1 parts 2,3-Ac(HO)C6H3O(CH2)3OC6H3(OH)Ac-3,2 (m. 184-5°), of which 6.9 parts in 15 parts (CO2Et)2 are added to a solution of Na 3 in EtOH 30 and C6H6 50 parts. The mixture is refluxed 20 hrs. and poured in Et20, the precipitate separated and dissolved in H2O, and the solution acidified to give a sticky precipitate which is boiled 10 min. with 50 parts EtOH, containing a catalytic amount of HCl. The mixture is cooled and the precipitate separated and worked up to give 7.8 parts I di-Et ester [A = (CH2)3, 5,5'-isomer] (IV), m. 182-3° (1:2 C6H6-EtOH). A suspension of 3 parts IV in 50 parts boiling EtOH is treated with 11.6 parts 1.015N aqueous NaOH and H2O added until a clear solution is obtained. Working up of the product gives the hydrate of the di-Na salt of I (A = (CH2)3, 5,5'isomer). Ethyl 7-hydroxychromone-2-carboxylate is heated with 0.5 equivalent 1,5-dibromopentane by heating in Me2CO in the presence of K2CO3 to give I diethyl ester [7,7'-isomer, A = (CH2)5], m. 148-50° (EtOH), [TABLE OMITTED] A

mixture of powdered Na 4.6, IV 7.44, and EtOAc 150 parts is refluxed 2.5 hrs. with stirring. The resulting orange solution is cooled and diluted with 400 parts Et20. The precipitate is extracted with H2O and the extract acidified. The precipitated oil is extracted with CHCl3 to give an oil which is refluxed 10 min. in EtOH and 0.5 part concentrated HCl. The solution is evaporated and the oily residue rubbed with Et20 to give 4.82 parts 1,5-bis(2-methylchromon-5-yloxy)pentane (V) (m. 140-3°). To a mixture of V 5 in dioxane 100, SeO2 6 parts is added and the mixture refluxed 6 hrs. to give VI. A mixture of PhCHO 1.5 and V 2.7 in EtOH 35 parts is added to EtONa (prepared from 0.294 part Na and 8 parts EtOH) and the solution refluxed 4 hrs. with stirring and left 16 hrs. at ambient temperature to give 1.55 parts 1,5-bis(2-styrylchromon-5vloxy) pentane (m. 217-20°), which is oxidized with KMnO4 to give VI. A mixture of 1,5-dibromopentane 5.7 in EtOH 80 is added to a solution of KOH 5.6 and hydroquinone 33 in EtOH 40 parts; the solution is refluxed 16 hrs., the EtOH evaporated, and 200 parts H2O added. The mixture is acidified and the precipitate separated and extracted with hot C6H6 to give 1.5diphenoxypentane, m. 110-12°, 5.8 parts of which is treated with a solution of 1.6 parts NaOH in 10 parts H2O. The H2O is evaporated and the precipitate stirred with 50 parts dioxane under reflux. Dropwise, 6.8 parts diethyl acetylenedicarboxylate is added, and the mixture refluxed 50 min. with stirring. After cooling and acidificaton, the solvent is evaporated to give VII. Ia and I prepared are tabulated in the 1st and 2nd tables.

AN 1967:500002 CAPLUS Full-text

DN 67:100002

OREF 67:18799a,18802a

TI Preparation of dichromonyl derivatives

PA Fisons Pharmaceuticals Ltd.

SO Neth. Appl., 53 pp.

CODEN: NAXXAN

DT Patent LA Dutch

FAN.CNT 1

FAN	PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
PI	NL 6603997	A	19660926	NL 1966-3997		19660325
				GB 1965-12626	A	19650325
				GB 1965-52414	A	19651209
				GB 1965-53744	A	19651217
	DE 1543579	B2	19800103	DE 1966-F48741		19660323
	DE 1543579	C3	19800828			
				GB 1965-12626	A	19650325
				GB 1965-52414	A	19651209
				GB 1965-53744	A	19651217
	DE 1792807	C2	19820819	DE 1967-1792807		19660323
				GB 1965-12626	A	19650325
				GB 1965-52414	A	19651209
				GB 1965-53744	A	19651217
	DK 134646	В	19761213	DK 1966-1562		19660325
	DK 134646	C	19770516			
				GB 1965-12626		19650325
				GB 1965-52414	A	19651209
				GB 1965-53744	A	19651217
	FI 56684	В	19791130	FI 1966-2334		19660907
	FI 56684	C	19800310			
	JP 53041662	В	19781106	JP 1970-111318		19701215
				GB 1965-12626	A	19650325
				GB 1965-52414	A	19651209
				GB 1965-53744	A	19651217
	JP 53041663	В	19781106	JP 1970-112555		19701217
				GB 1965-12626	Α	19650325

				GB	1965-52414	A	19651209
				GB	1965-53744	A	19651217
JP	53043492	В	19781120	JP	1971-1258		19710120
				GB	1965-12626	A	19650325
				GB	1965-52414	A	19651209
				GB	1965-53744	A	19651217
JP	53043493	В	19781120	JP	1971-12827		19710311
				GB	1965-12626	A	19650325
				GB	1965-52414	A	19651209
				GB	1965-53744	A	19651217
IN	137921	A1	19751011	IN	1975-CA734		19750414
				GB	1965-12626	A	19650325
				IN	1966-104449	A	19660322
FΙ	7903211	A	19791016	FI	1979-3211		19791016
				GB	1965-12626	A	19650325
					1966-2334	A	19660907

IT 15826-37-6F 16110-51-3P 16129-86-5F 16129-87-6F 16129-88-7F 16130-25-9F 16139-24-5F 16139-25-6P 16139-67-6F 16159-45-6F 16159-45-6F

RL: SPN (Synthetic preparation); PREP (Preparation)

(preparation of) RN 15826-37-6 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid,

5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[4-oxo-, sodium salt (1:2) (CA INDEX NAME)

RN 16110-51-3 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[4-oxo- (CA INDEX NAME)

RN 16129-86-5 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxytrimethylene)dioxy]bis[4-oxo-, calcium salt (1:1) (8CI) (CA INDEX NAME)

RN 16129-87-6 CAPLUS

4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxytrimethylene)dioxy]bis[4-oxo-, magnesium salt (1:1) (8CI) (CA INDEX NAME)

RN 16129-88-7 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxy-1,3-propanediyl)bis(oxy)]bis[4-oxo-, compd. with piperidine (1:2) (9C1) (CA INDEX NAME)

CM 1

CRN 16110-51-3 CMF C23 H16 O11

CM 2

CRN 110-89-4 CMF C5 H11 N RN 16130-23-7 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[7-methyl-4-oxo- (CA INDEX NAME)

RN 16130-25-9 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxytrimethylene)dioxy]bis[8-ethyl-4-oxo-, diethyl ester (8CI) (CA INDEX NAME)

RN 16139-24-5 CAPLUS

CN

4H-1-Benzopyran-2-carboxylic acid, 5-[3-[[2-(ethoxycarbonyl)-4-oxo-4H-1-benzopyran-5-yl]oxy]-2hydroxypropoxy]-8-ethyl-4-oxo-, ethyl ester (CA INDEX NAME)

RN 16139-25-6 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid, 5-[3-[(2-carboxy-4-oxo-4H-1-benzopyran-5-y1)oxy]-2-hydroxypropoxy]-8-ethyl-4-oxo (CA INDEX NAME)

PAGE 1-A

PAGE 2-A

RN 16139-67-6 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-ethoxyrrimethylene)dioxy]bis[4-oxo-, diethyl ester (8CI) (CA INDEX NAME)

RN 16146-53-5 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid,

5,5'-[(2-hydroxy-1,3-propanediy1)bis(oxy)]bis[8-ethy1-4-oxo- (CA INDEX NAME)

PAGE 1-A

PAGE 2-A

RN 16146-54-6 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid,

5,5'-[(2-ethoxy-1,3-propanediyl)bis(oxy)]bis[4-oxo-, disodium salt (9CI) (CA INDEX NAME)

RN 16150-45-1 CAPLUS

CN 4H-1-Benzopyran-2-carboxylic acid, 5,5'-[(2-hydroxy-1,3-propanediyl)bis(oxy)]bis[4-oxo-, 2,2'-diethyl ester (CA INDEX NAME)

$$\begin{array}{c|c} & & & & \\ & & & & \\ & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & \\ & & \\$$

OSC.G 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD (3 CITINGS)